

## SUMMARY OF PRODUCT CHARACTERISTICS

### 1. NAME OF MEDICINAL PRODUCT

Sodium Chloride 0.9 % w/v and Glucose 5% w/v Solution for Infusion BP

### 2. QUALITATIVE AND QUANTITATIVE COMPOSITION

Each 100ml contains Glucose (as anhydrous) 5g

S o d i u m    0.9g  
chloride:

mmol/l:            N a + :        Cl-: 154.  
                          154

MEq/l:             N a + :        Cl-: 154.  
                          154

pH: 3.5 – 6.5

For the full list of excipients: see section 6.1

### 3. PHARMACEUTICAL FORM

Solution for infusion

Clear solution, free from visible particles.

### 4. CLINICAL PARTICULARS

#### 4.1. Therapeutic Indications

Sodium Chloride 0.9 % w/v and Glucose 5% w/v solution is indicated for:

Treatment of sodium depletion, extracellular dehydration or hypovolaemia in cases where supply of water and carbohydrates is required due to restriction of the intake of fluids and electrolytes by normal routes of carbohydrate and fluid depletion

#### 4.2. Posology and method of administration

The choice of the specific sodium chloride and glucose concentration, dosage, volume, rate and duration of administration depends on the age, weight, clinical condition of the patient and concomitant therapy. It should be determined by a physician. For patients with electrolyte and glucose abnormalities and for pediatric patients, consult a physician experienced in intravenous fluid therapy.

Fluid balance, serum glucose, serum sodium and other electrolytes should be monitored before and during administration, especially in patients with increased non-osmotic vasopressin release (syndrome of inappropriate antidiuretic hormone secretion, SIADH) and in patients co-medicated with vasopressin agonist drugs due to the risk of hyponatraemia. Monitoring of serum sodium is particularly important for physiologically hypotonic fluids. Sodium Chloride 0.9 % w/v and Glucose 5% w/v solution may become extremely hypotonic after administration due to glucose metabolism in the body (see sections 4.4, 4.5 and 4.8).

Rapid correction of hyponatraemia and hypernatraemia is potentially dangerous (risk of serious neurologic complications);

Electrolyte supplementation may be indicated according to the clinical needs of the patient

**Adults, older patients and adolescents (age 12 years and over):**

The recommended dosage is: 500 ml to 3 L/24h

Administration rate:

The infusion rate is usually 40 ml/kg/24h and should not exceed the patient's glucose oxidation capacities in order to avoid hyper glycaemia. Therefore, the maximum acute administration rate is 5mg/kg/min.

**Paediatric population**

The dosage varies with weight:

- 0-10 kg body weight: 100 ml / kg / 24 h
- 10-20 kg body weight: 1000 ml + (50 ml/ kg over 10 kg) / 24h
- > 20 kg body weight: 1500 ml + (20 ml/ kg over 20 kg) / 24h.

The administration rate varies with weight:

- 0-10 kg body weight: 6-8 ml/kg/h
- 10-20 kg body weight: 4-6 ml/kg/h
- > 20 kg body weight: 2-4 ml/kg/h

The infusion rate should not exceed the patient's glucose oxidation capacities in order to avoid hyperglycaemia. Therefore, the maximum acute administration rate is 10-18 mg/kg/min depending on the total body mass.

For all patients, a gradual increase of flow rate should be considered when starting administration of glucose containing products.

**Method of administration**

The administration is performed by intravenous infusion.

Sodium chloride 0.9% w/v and Glucose 5% w/v solution is isotonic and hyperosmolar, due to the glucose content. It has an approximate osmolarity of 585 mOsmol/l.

**Precautions to be taken before manipulating or administering the product**

Parenteral drug products should be inspected visually for particulate matter and discoloration prior to administration. Do not administer unless the solution is clear and the seal is intact. Administer immediately following the insertion of infusion set. Do not remove unit from overwrap

until ready for use. The inner bag maintains the sterility of the product.

The solution should be administered with sterile equipment using an aseptic technique.

The equipment should be primed with the solution in order to prevent air entering the system.

Do not use plastic containers in series connections. Such use could result in air embolism due to residual air being drawn from the primary container before the administration of the fluid from the secondary container is completed. Pressurizing intravenous solutions contained in flexible plastic containers to increase flow rates can result in air embolism if the residual air in the container is not fully evacuated prior to administration. Use of a vented intravenous administration set with the vent in the open position could result in air embolism. Vented intravenous administration sets with the vent in the open position should not be used with flexible plastic containers

Additives may be introduced before or during infusion through the resealable medication port. When additive is used, verify tonicity prior to parenteral administration. Hyperosmolar solutions may cause venous irritation and phlebitis. Thus, any hyperosmolar solution is recommended to be administered through a large central vein, for rapid dilution of the hyperosmotic solution.

For further information on the product with additives, please see sections 6.2, 6.3 and 6.6.

#### **4.3. Contraindications**

The solution is contraindicated in patients presenting with:

- Known hypersensitivity to the product
- Extracellular hyperhydration or hypervolaemia
- Fluid and sodium retention
- Severe renal insufficiency (with oliguria/anuria)
- Uncompensated cardiac failure
- Hyponatraemia or hyperchloraemia
- General oedema and ascitic cirrhosis

Clinically significant hyperglycaemia. The solution is also contraindicated in case of uncompensated diabetes, other known glucose intolerances (such as metabolic stress situations), hyperosmolar coma or hyperlactatemia.

#### **4.4. Special warnings and precautions for use Hypokalaemia**

The infusion of Sodium chloride 0.9% w/v and Glucose 5% w/v solution may result in Hypokalaemia. Close clinical monitoring may be warranted in patients with or at risk for Hypokalaemia, for example:

- Persons with metabolic alkalosis
- Persons with thyrotoxic periodic paralysis. Administration of intravenous glucose has been associated in aggravating Hypokalaemia
- Persons with increased gastrointestinal losses (e.g., diarrhea, vomiting)
- Prolonged low potassium diet
- Persons with primary hyperaldosteronism
- Patients treated with medications that increase the risk of Hypokalaemia (e.g., diuretics,

beta-2agonists or insulin)

### **Sodium retention, fluid overload and oedema**

Sodium chloride 0.9% w/v and Glucose 5% w/v solution should be used with particular caution in

- Patients with metabolic acidosis
- Patients at risk of;
  - Hyponatraemia
  - Hyperchloraemia
  - Hypervolaemia
- Patients with conditions that may cause sodium retention, fluid overload and oedema (central and peripheral), such as
  - Primary hyperaldosteronism,
  - Secondary hyperaldosteronism associated with, for example,
    - hypertension,
    - congestive heart failure,
    - liver disease (including cirrhosis),
    - renal disease (including renal artery stenosis, nephrosclerosis)
  - Pre-eclampsia.

Patients taking medications that may increase the risk of sodium and fluid retention, such as corticosteroids

### ***Hyperosmolality, serum electrolytes and water imbalance***

Depending on the volume, rate of infusion, the patient's underlying clinical condition and capability to metabolize glucose, administration of Sodium chloride 0.9% w/v and Glucose 5% w/v solution can cause:

- Hyperosmolality, osmotic diuresis and dehydration
- Electrolyte disturbances such as
  - hyponatraemia (see "Sodium imbalance" below)
  - hypokalaemia (see above)
  - hypophosphataemia,
  - hypomagnesaemia,
- Acid-base imbalance
- Overhydration/hypervolaemia and, for example, congested states, including central (e.g. pulmonary congestion) and peripheral oedema.
- An increase in serum glucose concentration is associated with an increase in serum osmolality. Osmotic diuresis associated with hyperglycaemia can result in or contribute to the development of dehydration and in electrolyte losses.

### ***Electrolyte balance (see 'Sodium imbalance' below)***

Glucose intravenous infusions are usually isotonic solutions. In the body, however, glucose containing fluids can become extremely physiologically hypotonic due to rapid glucose metabolism (see section 4.2).

Depending on the tonicity of the solution, the volume and rate of infusion and depending on a patient's underlying clinical condition and capability to metabolize glucose, intravenous administration of glucose can cause electrolyte disturbances most importantly hypo- or hyperosmotic hyponatraemia.

### *Hyponatraemia:*

Patients with non-osmotic vasopressin release (e.g., in acute illness, pain, post-operative stress, infections, burns, and CNS diseases), patients with heart-, liver- and kidney diseases and patients exposed to vasopressin agonists (see section 4.5) are at particular risk of acute hyponatraemia upon infusion of hypotonic fluids.

Acute hyponatraemia can lead to acute hyponatraemic encephalopathy (brain oedema) characterized by headache, nausea, seizures, lethargy and vomiting. Patients with brain oedema are at particular risk of severe, irreversible and life-threatening brain injury.

Children, women in the fertile age and patients with reduced cerebral compliance (e.g. meningitis, intracranial bleeding, and cerebral contusion) are at particular risk of the severe and life-threatening brain swelling caused by acute hyponatraemia

Clinical evaluation and periodic laboratory determinations may be necessary to monitor changes in fluid balance, electrolyte concentrations, and acid-base balance during prolonged parenteral therapy or whenever the condition of the patient or the rate of administration warrants such evaluation.

### *Hyperglycaemia*

Rapid administration of glucose solutions may produce substantial hyperglycaemia and hyperosmolar syndrome. In order to avoid hyperglycaemia, the infusion rate should not exceed the patient's ability

to utilize glucose. To reduce the risk of hyperglycaemia-associated complications, the infusion rate must be adjusted and/or insulin administered if blood glucose levels exceed levels considered acceptable for the individual patient

Intravenous glucose should be administered with caution in patients with, for example:

- Impaired glucose tolerance (such as in diabetes mellitus, renal impairment, or in the presence of sepsis, trauma, or shock),
- Severe malnutrition (risk of precipitating a refeeding syndrome, see below),
- Thiamine deficiency, e.g., in patients with chronic alcoholism (risk of severe lactic acidosis due to impaired oxidative metabolism of pyruvate),
- Water and electrolyte disturbances that could be aggravated by increased glucose and/or free water load

Other groups of patients in whom Sodium chloride 0.9% w/v and Glucose 5% w/v solution should be used with caution include:

- Patients with ischemic stroke. Hyperglycaemia has been implicated in increasing cerebral ischemic brain damage and impairing recovery after acute ischemic strokes.
- Patients with severe traumatic brain injury (in particular during the first 24 hours following the trauma). Early hyperglycaemia has been associated with poor outcomes in patients with severe traumatic brain injury.
- Newborns (See Paediatric glycaemia-related issues).

Prolonged intravenous administration of glucose and associated hyperglycaemia may result in decreased rates of glucose-stimulated insulin secretion.

### *Hypersensitivity Reactions*

- Hypersensitivity/infusion reactions, including anaphylaxis, have been reported (see section 4.8).
- Stop the infusion immediately if signs or symptoms of hypersensitivity/infusion

reactions develop. Appropriate therapeutic countermeasures must be instituted as clinically indicated.

Solutions containing glucose should be used with caution in patients with known allergy to corn or corn products

#### *Refeeding syndrome*

Refeeding severely undernourished patients may result in the refeeding syndrome that is characterized by the shift of potassium, phosphorus, and magnesium intracellularly as the patient becomes anabolic. Thiamine deficiency and fluid retention may also develop. Careful monitoring and slowly increasing nutrient intake while avoiding overfeeding can prevent these complications

#### *Severe renal impairment*

Sodium chloride 0.9% w/v and Glucose 5% w/v solution should be administered with particular caution to patients at risk of (severe) renal impairment. In such patients, administration may result in sodium retention and/or fluid overload.

#### *Paediatric use*

The infusion rate and volume depend on the age, weight, clinical and metabolic conditions of the patient, concomitant therapy, and should be determined by a physician experienced in paediatric intravenous fluid therapy.

#### *Paediatric glycaemia-related issues*

Newborns, especially those born premature and with low birth weight, are at increased risk of developing hypo- or hyperglycaemia. Close monitoring during treatment with intravenous glucose solutions is needed to ensure adequate glycaemic control, in order to avoid potential long term adverse effects.

- Hypoglycaemia in the newborn can cause, e.g., prolonged seizures, coma, and cerebral injury
- Hyperglycaemia has been associated with cerebral injury, including intraventricular haemorrhage, late onset bacterial and fungal infection, retinopathy of prematurity, necrotizing enterocolitis, increased oxygen requirements, prolonged length of hospital stay, and death.

#### *Paediatric hyponatraemia-related issues*

Children (including neonates and older children) are at increased risk of developing hyponatraemia as well as for developing hyponatraemic encephalopathy.

- Hyponatraemia can lead to headache, nausea, seizures, lethargy, coma, cerebral oedema and death; therefore, acute symptomatic hyponatraemic encephalopathy is considered a medical emergency.
- Plasma electrolyte concentrations should be closely monitored in the paediatric population
- Rapid correction of hyponatraemia is potentially dangerous (risk of serious neurologic complications). Dosage, rate, and duration of administration should be determined by a physician experienced in paediatric intravenous fluid therapy

#### *Blood*

Sodium chloride 0.9% w/v and Glucose 5% w/v solution should not be administered simultaneously with blood through the same administration set because of the possibility of pseudoagglutination or haemolysis.

### *Geriatric use*

When selecting the volume/rate of infusion for a geriatric patient, consider that geriatric patients are

#### **4.5. Interaction with other medicinal products and other forms of interaction**

Both the glycaemic effects of Glucose 5% and its effects on water and electrolyte balance should be taken into account when using Glucose 5% in patients treated with other substances that affect glycaemic control, or fluid and/or electrolyte balance.

Concomitant administration of catecholamines and steroids decreases the glucose up-take.

#### *Drugs leading to an increased vasopressin effect*

The below listed drugs increase the vasopressin effect, leading to reduced renal electrolyte free water excretion and increase the risk of hospital acquired hyponatraemia following inappropriately balanced treatment with IV fluids (see sections 4.2, 4.4 and 4.8).

- Drugs stimulating vasopressin release, e.g.: Chlorpropamide, clofibrate, carbamazepine, vincristine, selective serotonin reuptake inhibitors, 3,4-methylenedioxy-N-methamphetamine, ifosfamide, antipsychotics, narcotics
- Drugs potentiating vasopressin action, e.g.: Chlorpropamide, NSAIDs, cyclophosphamide
- Vasopressin analogues, e.g.: Desmopressin, oxytocin, terlipressin

Other medicinal products increasing the risk of hyponatraemia also include diuretics in general and antiepileptics such as oxcarbazepine.

Caution is advised in patients treated with

- lithium. Renal sodium and lithium clearance may be increased during administration and can result in decreased lithium levels.
- corticosteroids, which are associated with the retention of sodium and water (with oedema and hypertension).
- diuretics, beta-2 agonists or insulin, whom increase the risk of Hypokalaemia

#### **4.6. Fertility, pregnancy and lactation**

##### ***Pregnancy***

Intrapartum maternal intravenous glucose infusion may result in foetal hyperglycaemia and metabolic acidosis as well as rebound neonatal hypoglycaemia due to foetal insulin production.

##### ***Fertility***

There is no information on the effects of Sodium chloride 0.9% w/v and Glucose 5% w/v solution on fertility

##### ***Lactation***

Sodium chloride 0.9% w/v and Glucose 5% w/v solution can be used during breast-feeding.

The potential risks and benefits for each specific patient should be carefully considered before administration.

#### **4.7. Effects on ability to drive and use machines**

There is no information on the effects of Sodium chloride 0.9% w/v and Glucose 5% w/v solution

on the ability to operate an automobile or other heavy machinery.

Sodium chloride 0.9% w/v and Glucose 5% w/v solution should be administered with special caution for pregnant women during labour particularly if administered in combination with oxytocin due to the risk of hyponatraemia (see section 4.4, 4.5 and 4.8).

#### 4.8. Undesirable effects

The adverse drug reactions listed in this section are given following the recommended frequency convention: very common ( $\geq 1/10$ ); common ( $\geq 1/100$  to  $< 1/10$ ); uncommon ( $\geq 1/1000$  to  $< 1/100$ ); rare ( $\geq 1/10,000$  to  $< 1/1,000$ ); very rare ( $< 1/10,000$ ); and not known (cannot be estimated from the available data).

System Organ Class (SOC)	Adverse reactions (Preferred Term)	Frequency
Immune system disorders	Anaphylactic reaction* Hypersensitivity*	Not known
Metabolism and nutrition disorders	Hyponatremia Hyperglycaemia Hospital acquired hyponatraemia**	Not known
Nervous system disorders	Hyponatraemic encephalopathy**	Not known
Skin and subcutaneous tissue disorders	Rash Pruritus	Not known
Vascular disorders	Phlebitis	Not known
Renal and urinary disorders	Polyuria	Not known
General disorders and administration site conditions	Infusion site irritation including Chills Pyrexia Infusion site infection	Not known

\*Potential manifestation in patients with allergy to corn, see section 4.4.

\*\* Hospital acquired hyponatraemia may cause irreversible brain injury and death due to development of acute hyponatraemic encephalopathy (see sections 4.2 and 4.4).

Other adverse reactions reported with isotonic saline and glucose infusions include:

- Hyponatraemia, which may be symptomatic
- Hyperchloraemic acidosis

Adverse reactions may be associated to the medicinal product(s) added to the solution; the nature of the additive will determine the likelihood of any other adverse reactions.

#### Reporting of suspected adverse reactions



It's crucial to report any possible adverse reactions; It enables ongoing evaluation of the pharmaceutical product's benefit/risk ratio. Health care professionals are asked to report any suspected adverse reactions via the **TMDA** yellowforms.

#### **4.9.Overdose**

Excess administration of Sodium chloride 0.9% w/v and Glucose 5% w/v solution can cause:

- Hyperglycaemia, adverse effects on water and electrolyte balance and corresponding complications. For example, severe hyperglycaemia and severe dilutional hyponatraemia and their complications, can be fatal.
- Hyponatraemia (which can lead to CNS manifestations, including seizures, coma, cerebral oedema and death).
- Hypernatraemia especially in patients with renal impairment.
- Fluid overload (which can lead to central and/or peripheral oedema).
- See also sections 4.4 and 4.8

A clinically significant overdose of Sodium chloride 0.9% w/v and Glucose 5% w/v solution may, therefore, constitute a medical emergency

When assessing an overdose, any additives in the solution must also be considered.

Interventions include discontinuation of Sodium chloride 0.9% w/v and Glucose 5% w/v solution administration, dose reduction, administration of insulin and other measures as indicated for the specific clinical constellation.

### **5. PHARMACOLOGICAL PROPERTIES**

#### **5.1. Pharmacodynamic properties**

Pharmacotherapeutic group "Electrolytes with Carbohydrates",

ATC code: "B05BB02".

Sodium Chloride 0.9% w/v and Glucose 5% w/v is an isotonic and hyperosmolar solution of sodium chloride and glucose.

The pharmacodynamic properties of this solution are those of its components (glucose, sodium and chloride).

Ions, such as sodium, circulate through the cell membrane, using various mechanisms of transport, among which is the sodium pump ( $\text{Na}^+/\text{K}^+$ -ATPase). Sodium plays an important role in neurotransmission and cardiac electrophysiology, and also in renal metabolism.

Chloride is mainly an extracellular anion. Intracellular chloride is in high concentration in red blood cells and gastric mucosa. Reabsorption of chloride follows reabsorption of sodium.

Glucose is the principal source of energy in cellular metabolism. The glucose in this solution provides a caloric intake of 200 kcal/l.

#### **5.2. Pharmacokinetic properties**

The pharmacokinetic properties of this solution are those of its components (glucose, sodium and chloride).

After injection of radio sodium ( $^{24}\text{Na}$ ), the half-life is 11 to 13 days for 99% of the injected Na and one year for the remaining 1%. The distribution varies according to tissues: it is fast in

muscles, liver, kidney, cartilage and skin; it is slow in erythrocytes and neurons; it is very slow in the bone. Sodium is predominantly excreted by the kidneys, but (as described earlier) there is extensive renal reabsorption. Small amounts of sodium are lost in the faeces and sweat.

The two main metabolic pathways of glucose are gluconeogenesis (energy storage) and glycogenolysis (energy release). Glucose metabolism is regulated by insulin.

### **5.3.Pre-clinical safety data**

Preclinical safety data of this solution for infusion in animals are not relevant since its constituents are physiological components of animal and human plasma.

Toxic effects are not to be expected under the condition of clinical application. The safety of potential additives should be considered separately.

The safety of the additive should be considered separately.

## **6. PHARMACEUTICAL EXCIPIENTS**

### **6.1.List of excipients**

Water For Injections

### **6.2.Incompatibilities**

Incompatibility of the medicinal product to be added with the solution in the Viaflo container must be assessed before addition. In the absence of compatibility studies, this solution must not be mixed with other medicinal products.

The instructions for use of the medicinal product to be added must be consulted.

Before adding a drug, verify it is soluble and stable in water at the pH of Sodium Chloride 0.9% w/v and Glucose 5% w/v solution (see section 3).

As guidance, the following medications are incompatible with the Sodium Chloride 0.9 % w/v & Glucose 5% w/v solution (non-exhaustive listing):

- Ampicillin sodium
- Mitomycin
- Amphotericin B
- Erythromycin lactobionate

Those additives known to be incompatible should not be used.

Because of the presence of glucose, Sodium chloride 0.9% w/v and Glucose 5% w/v solution should not be administered simultaneously with blood through the same administration set because of the possibility of pseudoagglutination or haemolysis

### **6.3.Shelf life**

12 months

### **6.4.Special precautions for storage**

Do not store above 30°C. Protect from Sunlight  
For Intravenous use only

#### **6.5. Nature and contents of container**

Polypropylene bottles with an infusion site, or injection site for addition of medicinal products;  
The bottles are individually overwrapped in transparent Biaxially Oriented Polypropylene Film  
Bottles contain 500ml solution.

#### **6.6. Special precautions for disposal and other handling**

Discard any unused portion.

Do not store solutions containing additives. Do not reconnect partially used bags.

Do not remove unit from overwrap until ready for use. The inner bag maintains the sterility of the product.

When introducing additives to the infusion, aseptic techniques must be used. Mix the solution thoroughly when additives have been introduced.

#### **7. MARKET AUTHORIZATION HOLDER**

Alfa Pharmaceuticals Limited  
Plot No. 14 Mbagala Industrial Area  
P.O BOX 5804 Dar es salaam

#### **8. MARKETING AUTHORISATION NUMBER(S)**

TAN 22 HM 0468

#### **9. DATE OF FIRST AUTHORIZATION/ RENEWAL OF AUTHORIZATION**

31<sup>st</sup> October, 2022

#### **10. DATE OF REVISION OF TEXT**