

Short Stature

Jamal Al Jubeh, MD, FAAP
Consultant Pediatric Endocrinologist
Chairman of Pediatrics
SKMC, Abu Dhabi, UAE

Objectives

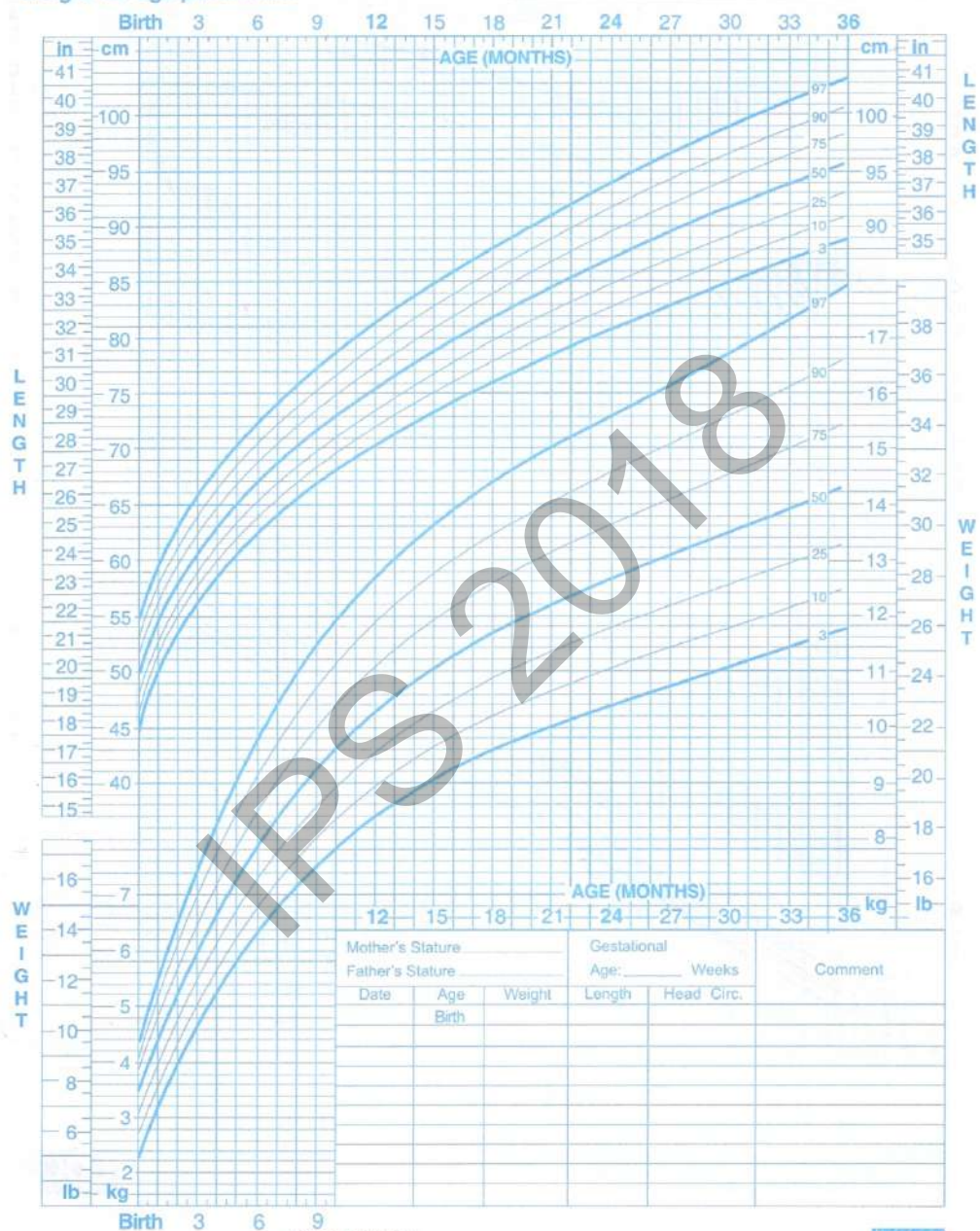
- Normal and abnormal growth patterns.
- Short stature and growth failure.
- Causes of growth failure.
- Evaluation process.



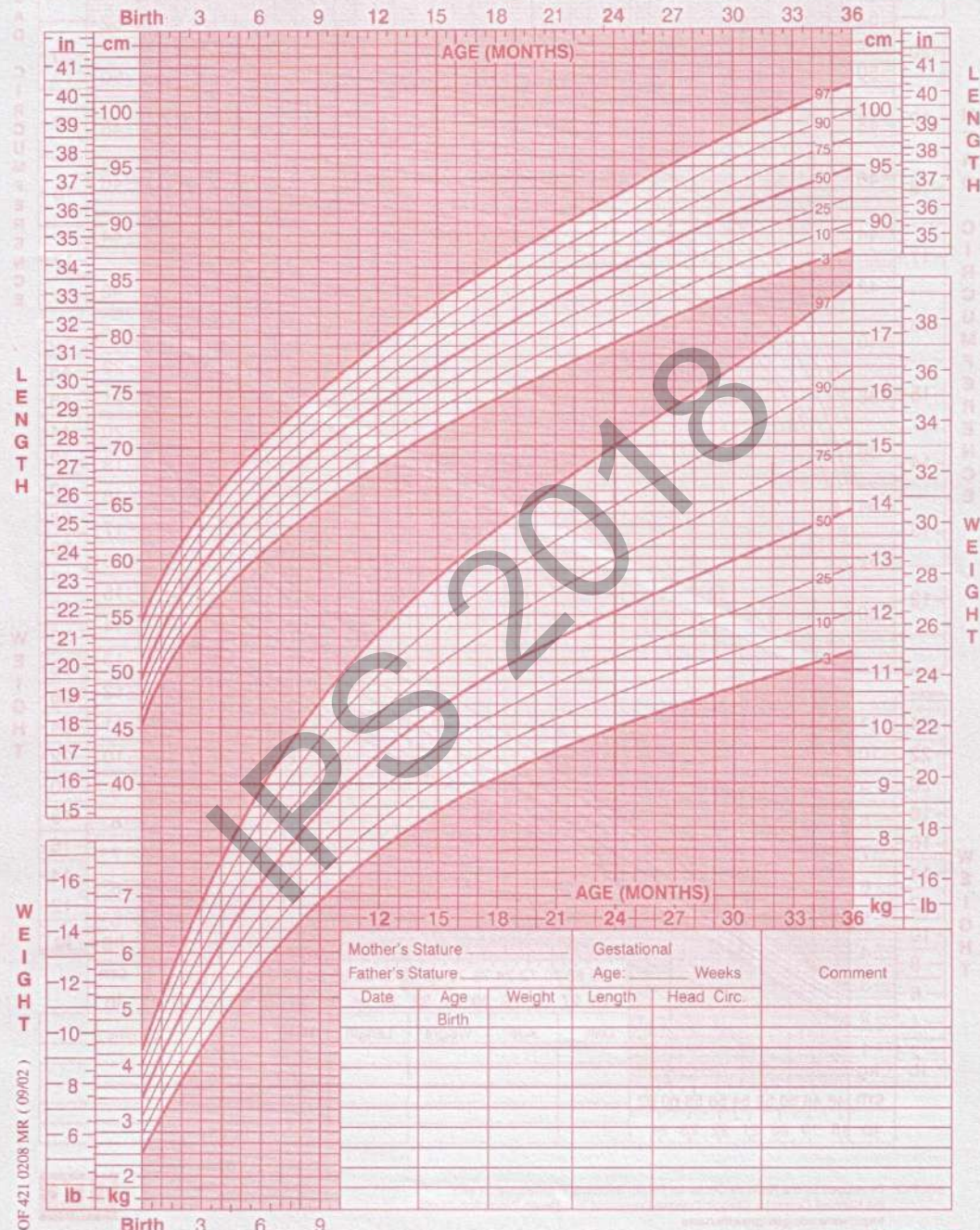
What is normal growth?

IPS 2018

Birth to 36 months: Boys
 Length-for-age and
 Weight-for-age percentiles



Length-for-age and Weight-for-age percentiles



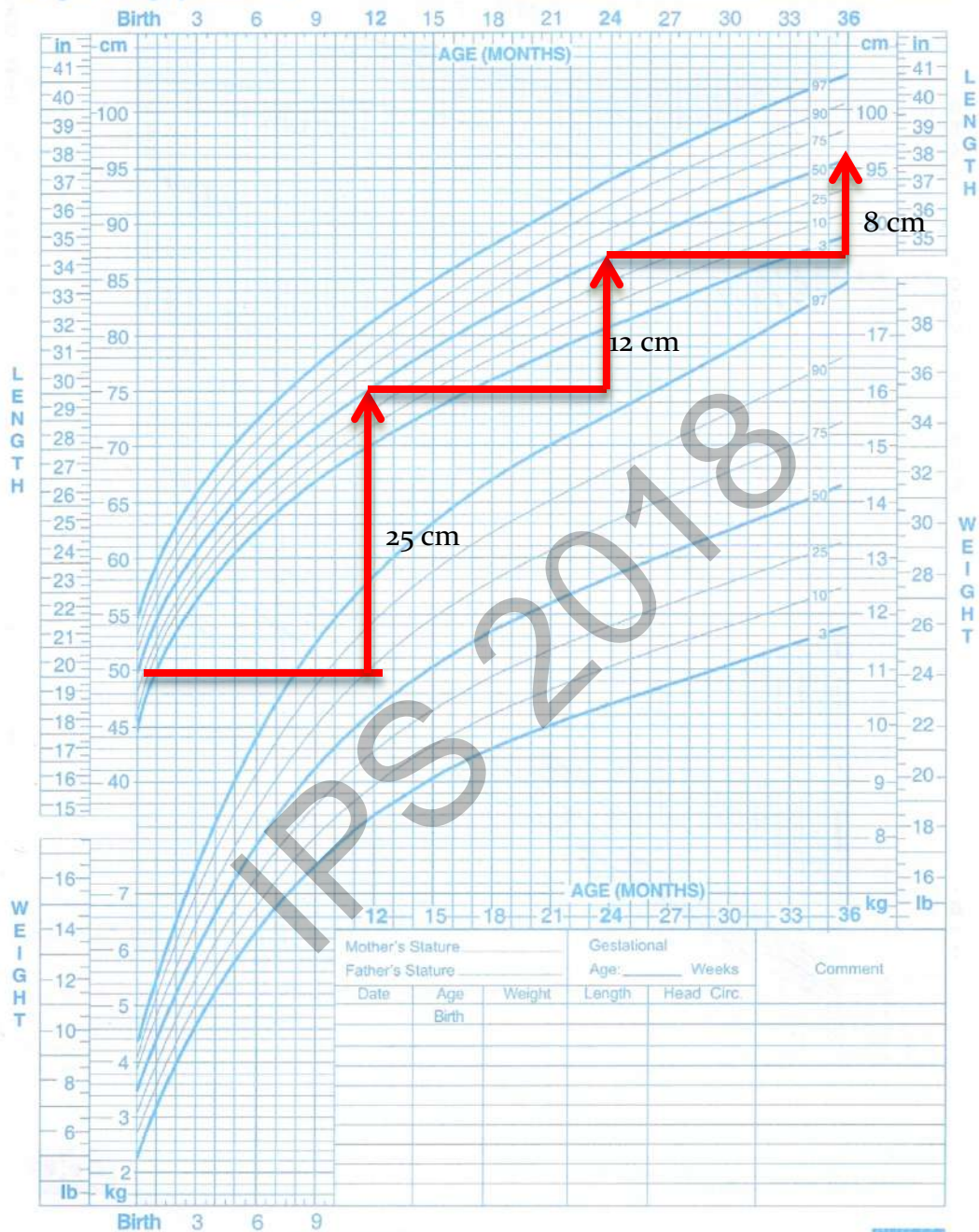
Mother's Stature _____			Gestational Age: _____ Weeks			Comment
Father's Stature _____						
Date	Age Birth	Weight	Length	Head Circ.		

OF 421 0208 MR (09/02)

Revised April 20, 2001.



Weight-for-age percentiles



Growth Velocity curves

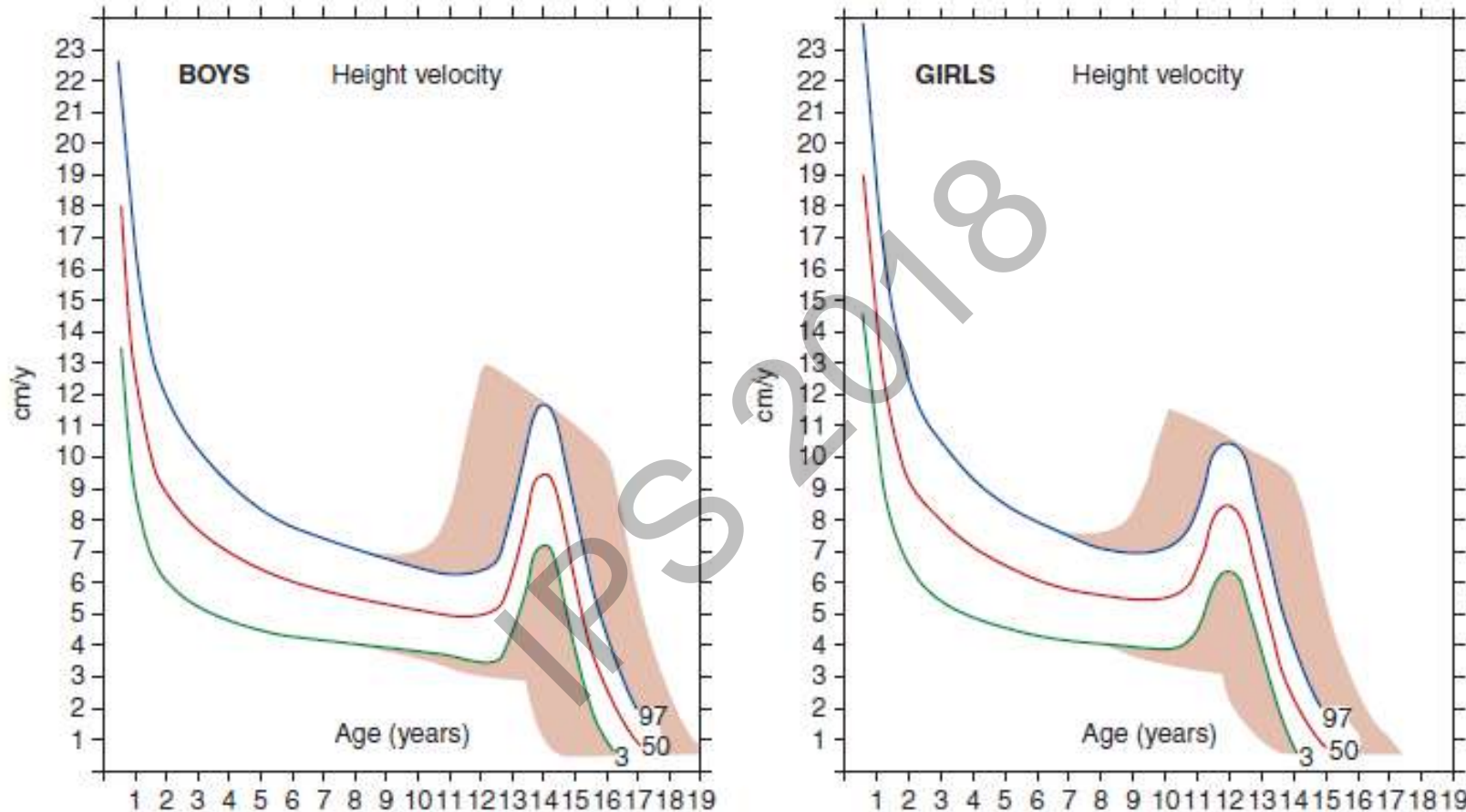


FIGURE 2-2 ■ Ranges of linear growth velocities in males and females. (Modified from charts prepared by Tanner and Whitehouse, 1976, and reproduced with permission of Tanner JM and Castlemead Publications, Ward's Publishing Services, Herts, UK.)

Table 1. Normal Growth Velocity at Various Life Stages

<i>Life stage</i>	<i>Growth velocity per year</i>
In utero	60 to 100 cm (24 to 40 in)
First year	23 to 27 cm (9 to 11 in)
Second year	10 to 14 cm (4 to 6 in)
Fourth year	6 to 7 cm (2 to 3 in)
Prepubertal nadir	5 to 5.5 cm (2 to 2.2 in)
Pubertal growth spurt	Girls: 8 to 12 cm (3 to 5 in) Boys: 10 to 14 cm (4 to 6 in)

Upper to Lower Extremity Ratios in Boys and Girls

Age (Years)	Boys	Girls
	U/L Ratio	U/L Ratio
Birth	1.70	1.70
1/2	1.62	1.60
1	1.54	1.52
1 1/2	1.50	1.46
2	1.42	1.41
2 1/2	1.37	1.34
3	1.35	1.30
3 1/2	1.30	1.27
4	1.24	1.22
4 1/2	1.22	1.19
5	1.19	1.15
6	1.12	1.10
7	1.07	1.06
8	1.03	1.02
9	1.02	1.01
10	0.99	1.00
11	0.95	0.90
12	0.98	0.99
13	0.97	1.00
14	0.97	1.01
15	0.95	1.01
16	0.99	1.01
17	0.99	1.01

Adapted from Wilkins L. *The Diagnosis and Treatment of Endocrine Disorders in Childhood and Adolescence*. IL: Springfield, Charles C. Thomas, Publisher, 1957.

How to measure length and height?

IPS 2018

Measurement of length

A



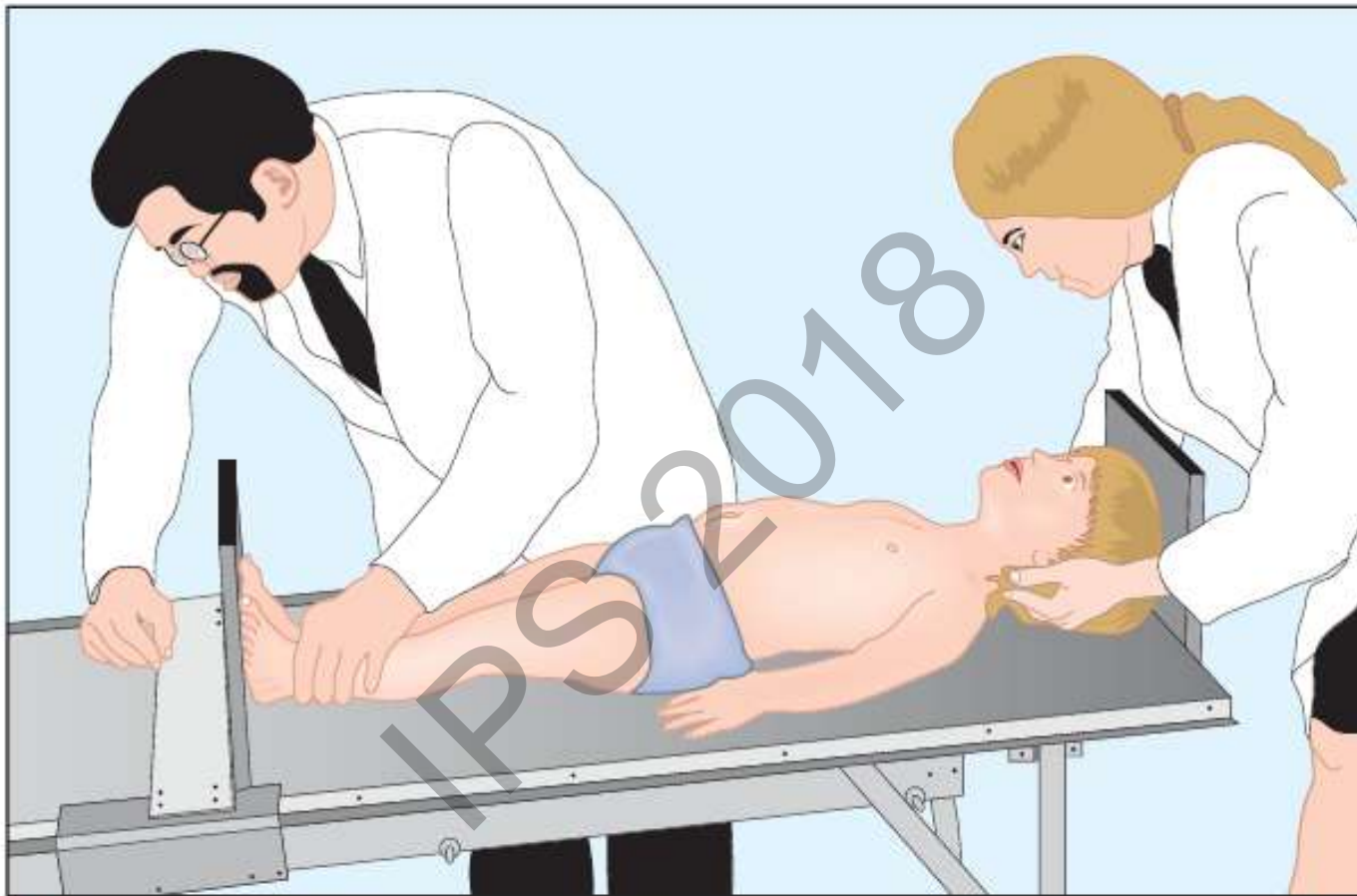
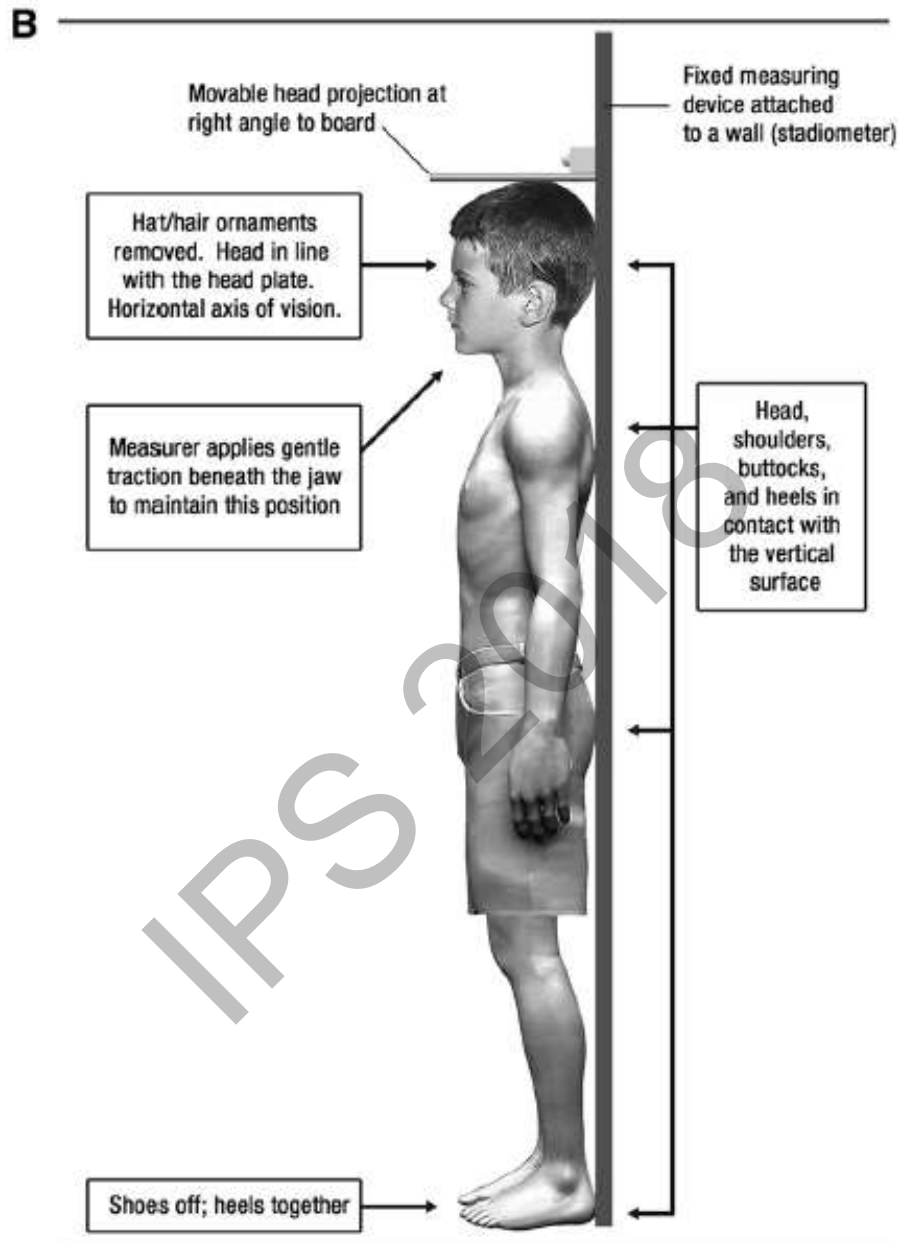
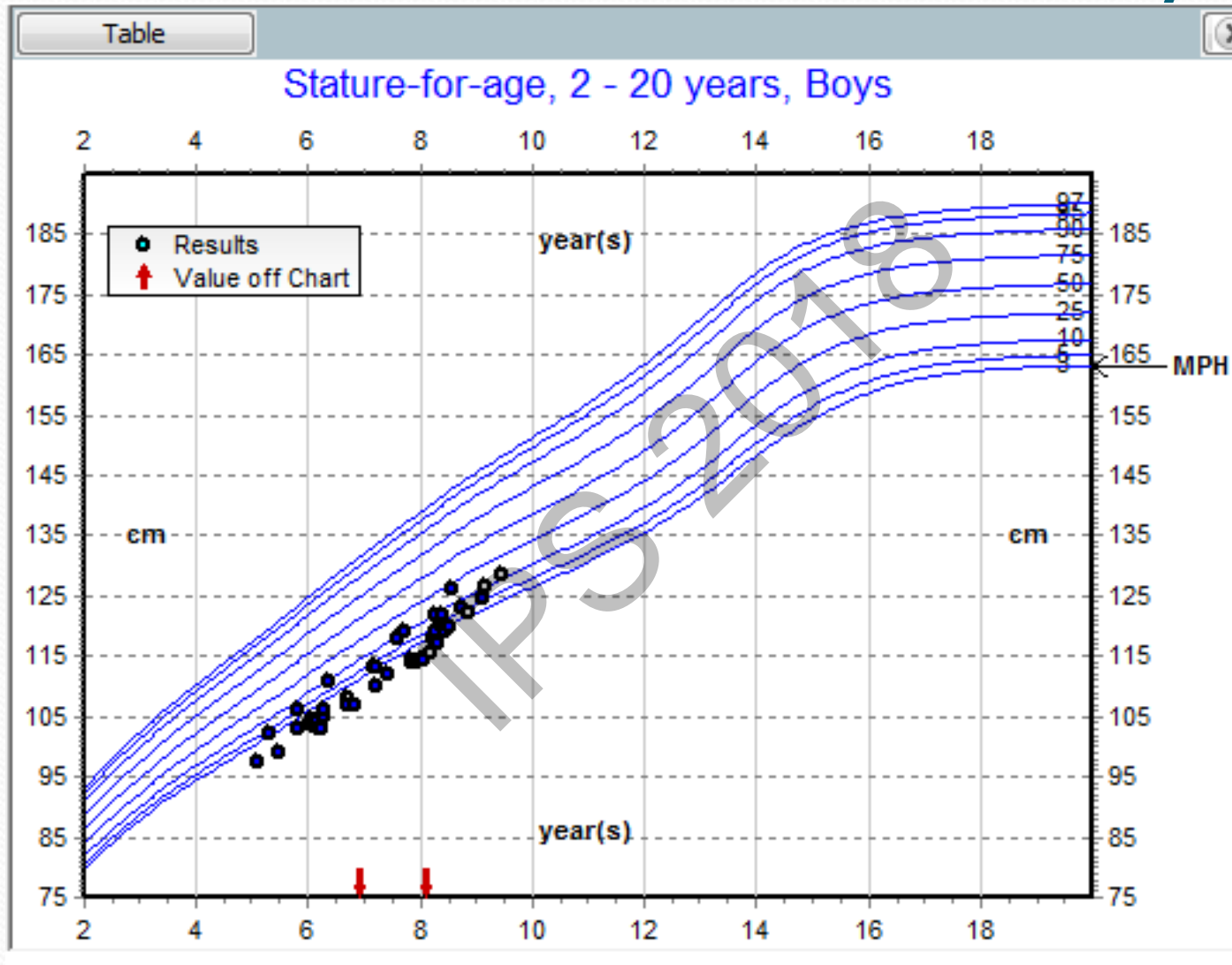


FIGURE 2-3 ■ Measurement of children less than 2 years of age should be obtained in the “Frankfurt plane” which places children in the supine position in full extension and other canthus of the eyes and the external auditory meatus perpendicular to the long axis of the trunk.

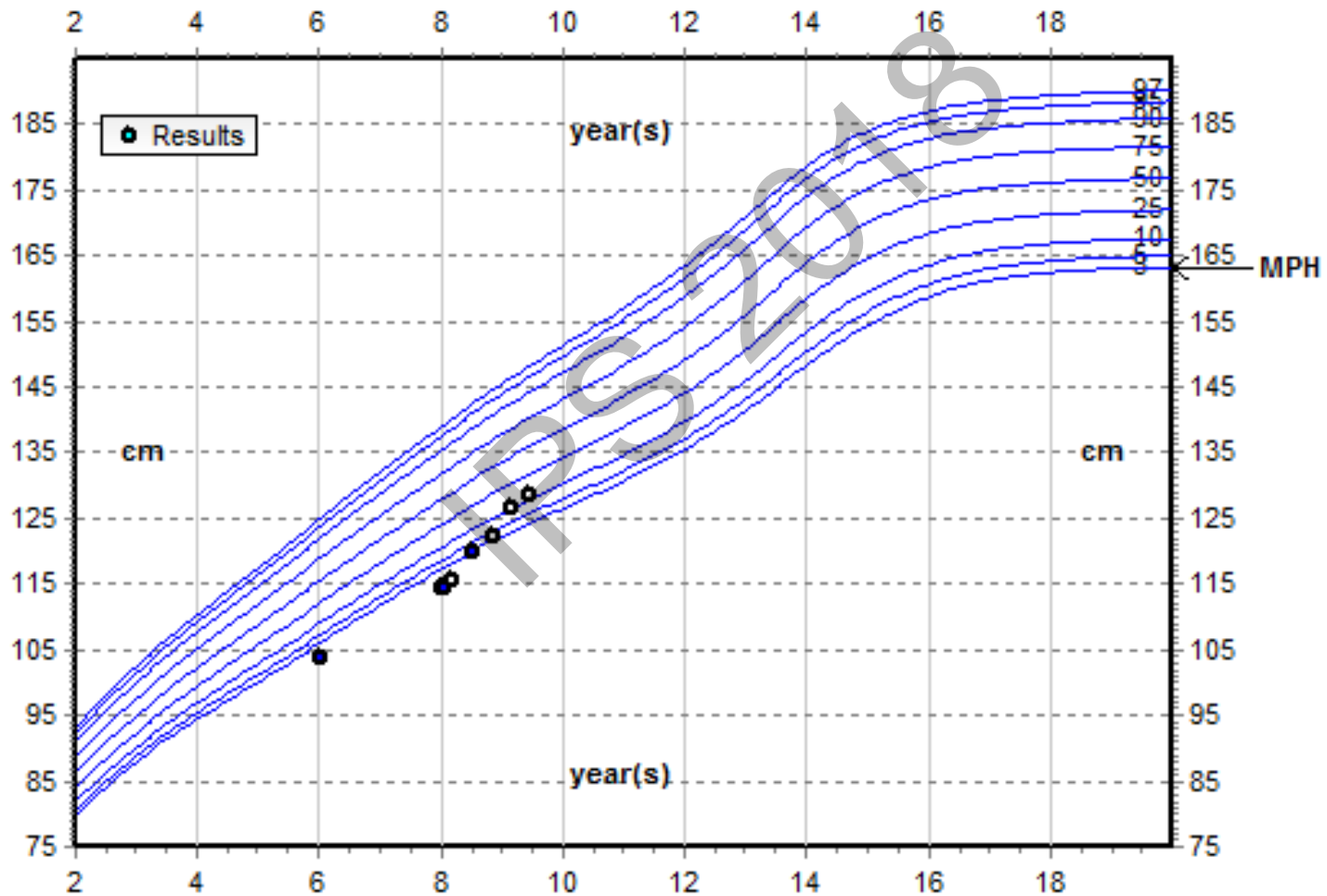



Measurement accuracy



<input type="button" value="Chart"/> <input type="button" value="Calculate GV"/> <input type="button" value="Plot All"/>							
Stature-for-age, 2 - 20 years, Boys							
Date	Age	Value	Centile	z-score/SD	GV Calculation	Medical Service	Plot
17/03/2013	8 years	(c) 115.40 cm	0.78	-2.42	<input type="checkbox"/>	Paediatrics-Endocrinology	<input checked="" type="checkbox"/>
08/04/2013	8 years	118.00 cm	2.58	-1.95	<input type="checkbox"/>	Family Practice	<input type="checkbox"/>
15/04/2013	8 years	119.00 cm	3.26	-1.84	<input type="checkbox"/>	Family Practice	<input type="checkbox"/>
16/04/2013	8 years	122.00 cm	9.48	-1.31	<input type="checkbox"/>	Family Practice	<input type="checkbox"/>
09/05/2013	8 years	117.00 cm	1.38	-2.20	<input type="checkbox"/>	Med-Dermatology	<input type="checkbox"/>
05/06/2013	8 years	122.00 cm	8.25	-1.39	<input type="checkbox"/>	Med-Dermatology	<input type="checkbox"/>
08/06/2013	8 years	120.00 cm	4.08	-1.74	<input type="checkbox"/>	Pharmacy	<input type="checkbox"/>
18/06/2013	8 years	119.00 cm	2.31	-1.99	<input type="checkbox"/>	Family Practice	<input type="checkbox"/>
14/07/2013	8 years	120.00 cm	2.95	-1.89	<input type="checkbox"/>	Paediatrics-Endocrinology	<input checked="" type="checkbox"/>
27/07/2013	8 years	126.00 cm	19.87	-0.85	<input type="checkbox"/>	Family Practice	<input type="checkbox"/>
03/10/2013	8 years	123.00 cm	6.58	-1.51	<input type="checkbox"/>	Family Practice	<input type="checkbox"/>
24/11/2013	8 years	(c) 122.40 cm	4.01	-1.75	<input type="checkbox"/>	Paediatrics-Endocrinology	<input checked="" type="checkbox"/>
17/02/2014	9 years	124.50 cm	5.59	-1.59	<input type="checkbox"/>	Paediatrics-General	<input type="checkbox"/>
09/03/2014	9 years	(c) 126.60 cm	10.76	-1.24	<input type="checkbox"/>	Paediatrics-Endocrinology	<input checked="" type="checkbox"/>
22/06/2014	9 years	(c) 128.40 cm	11.51	-1.20	<input type="checkbox"/>	Paediatrics-Endocrinology	<input checked="" type="checkbox"/>

Stature-for-age, 2 - 20 years, Boys





Short Stature: Definition, causes and approach

IPS 2018

Short Stature

- Height more than 2 SD below mean for the same age and sex compared to genetically relevant population (< 2.3 %)

IPS 2018

Growth evaluation

- Accurate measurement of child's height and weight.
- Accurate plotting on appropriate growth chart.
- Assessment of longitudinal growth (growth velocity):
Accurate heights measured at 6 - 12 month interval
- Measuring parent's height.

Familial (genetic) Short stature

- Parents are short
- Child is growing within target height range.
- Child's bone age = Chronological age

Midparental height (Boys)

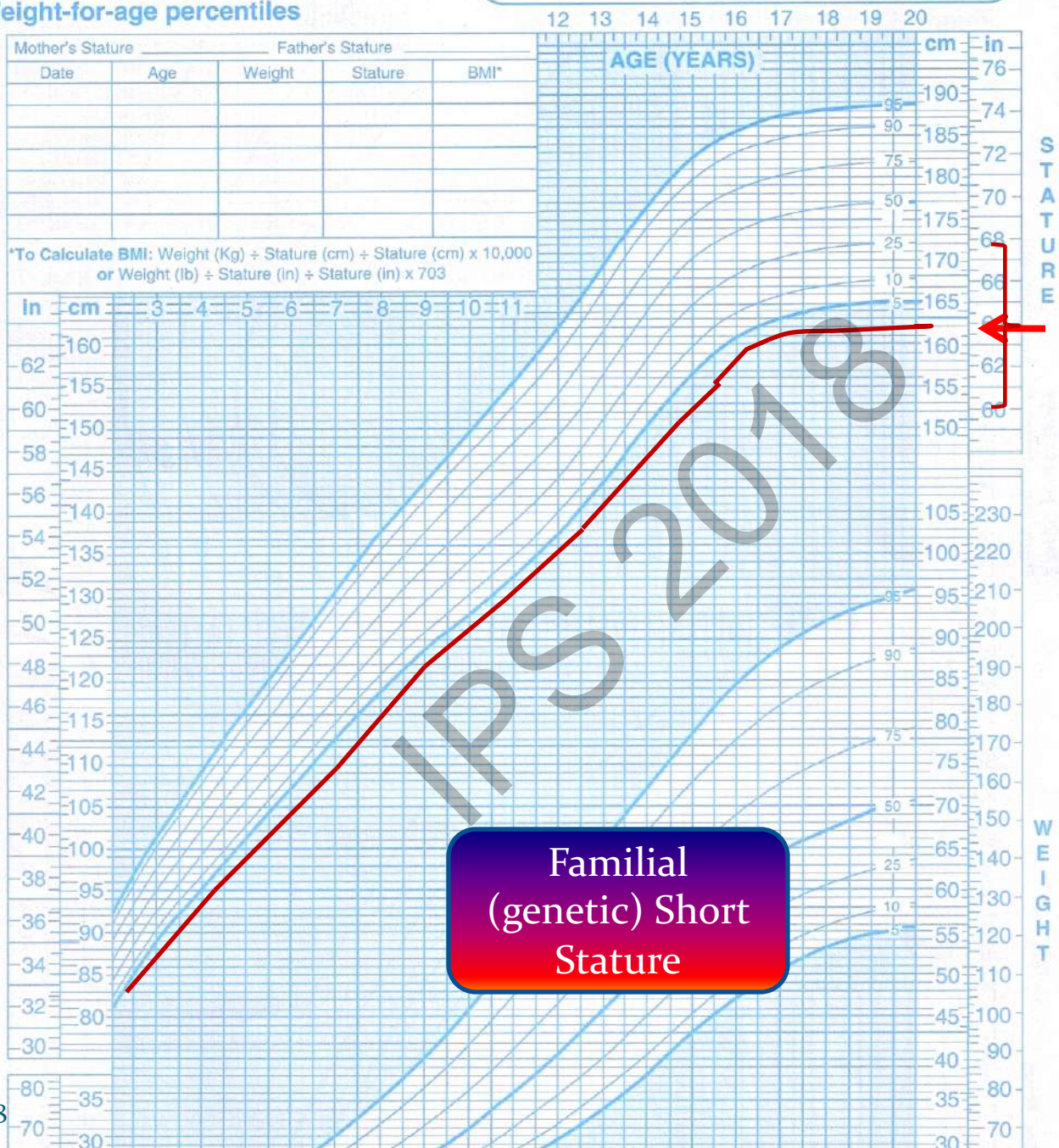
- F: Father's height (cm)
M: Mother's height (cm)
- Midparental height in cm for boys
- $= (F + M + 13) / 2$
- $= (F + M) / 2 + 6.5$

Midparental height (Girls)

- F: Father's height (cm)
M: Mother's height (cm)
- Midparental height in cm for Girls
- $$= (F + M - 13) / 2$$
- $$= (F + M) / 2 - 6.5$$

2 to 20 years: Boys
Stature-for-age and
Weight-for-age percentiles

Date of Admission OPD IPD



Mother's Stature		Father's Stature		
Date	Age	Weight	Stature	BMI*

*To Calculate BMI: Weight (Kg) ÷ Stature (cm) ÷ Stature (cm) x 10,000
or Weight (lb) ÷ Stature (in) ÷ Stature (in) x 703

**Familial
(genetic) Short
Stature**

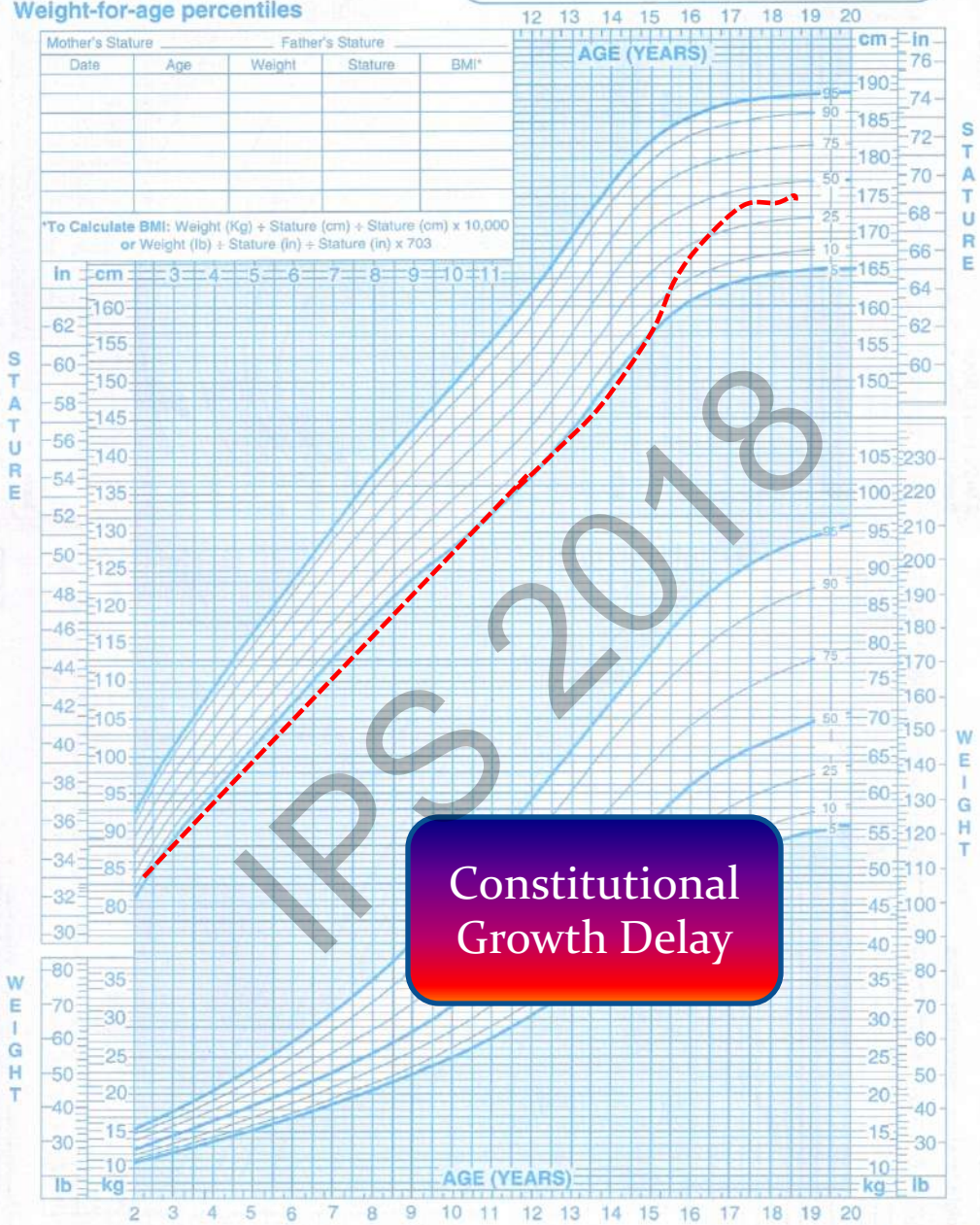
Constitutional Delay of Growth and Puberty (CDGP)

- Normal growth velocity.
- Delayed bone age.
- Delayed puberty (Often runs in the family)
- Normal adult (final) height

2 to 20 years: Boys
 Stature-for-age and
 Weight-for-age percentiles

Date of Admission :

OPD
 IPD



Mother's Stature		Father's Stature		
Date	Age	Weight	Stature	BMI*

*To Calculate BMI: Weight (Kg) ÷ Stature (cm) ÷ Stature (cm) x 10,000
 or Weight (lb) ÷ Stature (in) ÷ Stature (in) x 703

In	cm	3	4	5	6	7	8	9	10	11
62	160									
60	155									
58	150									
56	145									
54	140									
52	135									
50	130									
48	125									
46	120									
44	115									
42	110									
40	105									
38	100									
36	95									
34	90									
32	85									
30	80									

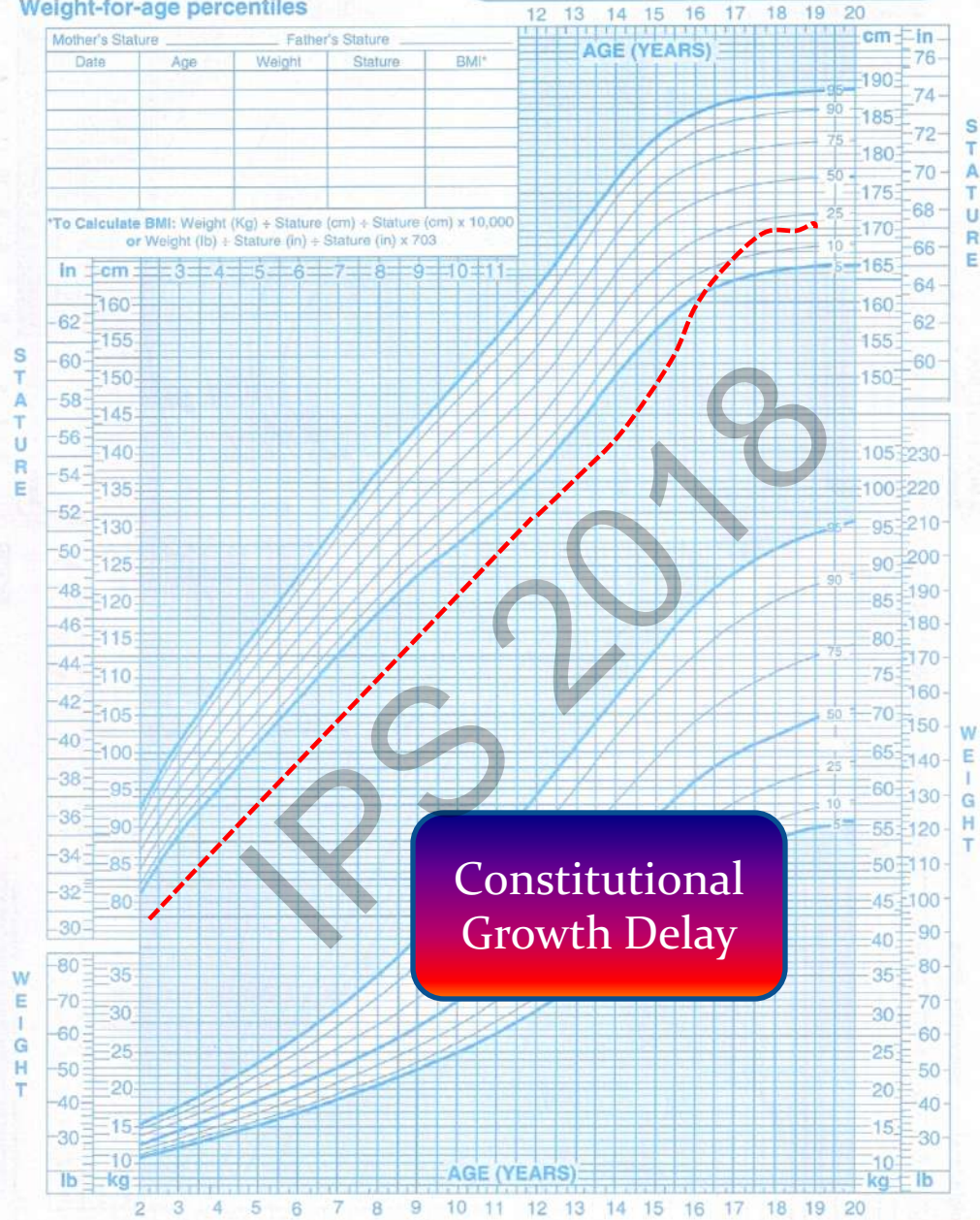
Constitutional Growth Delay



2 to 20 years: Boys
 Stature-for-age and
 Weight-for-age percentiles

Date of Admission :

OPD
 IPD



Constitutional Growth Delay





What are the causes of short stature?

Genes

Nutrition

Syndromes

Causes of
Short Stature

Systemic
Disease

Skeletal
Dysplasia

Hormones

SGA

GI

- Celiac Disease
- Malabsorption
- IBD

Renal

- CKD
- Renal Tubular Acidosis

Systemic Diseases

Hem / ONC

- Thalassemia
- SCD
- Malignancy

Pulmonary

- Cystic Fibrosis
- CLD

Metabolic

- Organic Acidemia
- GSD

Cardiac

Congenital
Heart disease

Endocrine

- Hypothyroidism
- Isolated growth hormone deficiency
- Hypopituitarism
- GH insensitivity (Laron dwarfism)
- Cushing syndrome
- Poorly controlled type 1 diabetes mellitus - Mauriac syndrome
- Rickets
- Idiopathic short stature

Evaluation of Short Stature

Should be based on findings from
History and Physical Exam

Evaluation of Short stature

- CBC, ESR
- Electrolytes & Renal Panel, Ca, Phos.
- Thyroid (TSH, FT₄)
- Celiac Screen:
 - (Tissue transglutaminase antibodies).
- Chromosomal analysis (girls).
- IGF-1, IGF-BP₃
- Bone Age X ray.

Differential diagnosis of low circulating IGF-I concentration

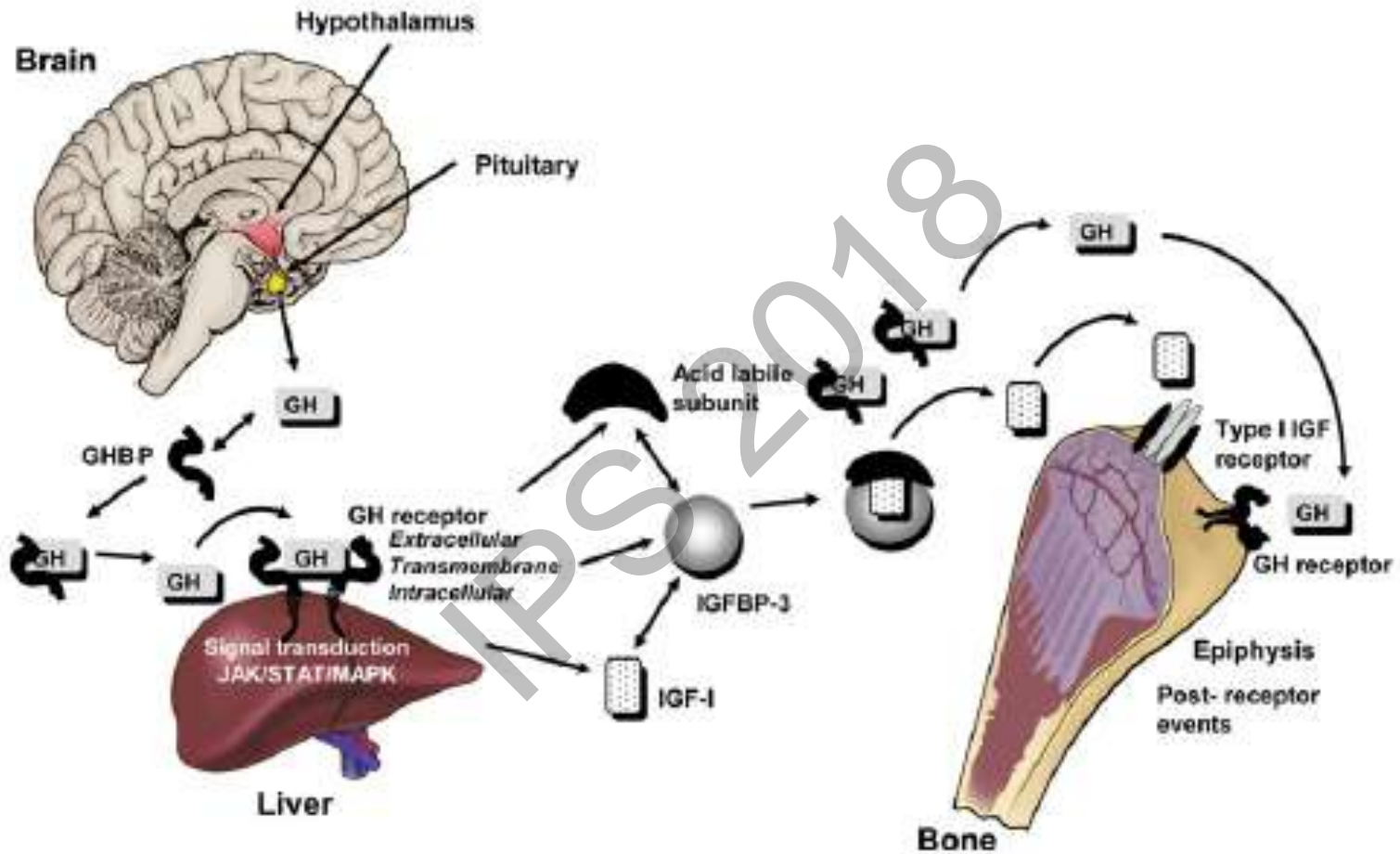
- GH deficiency
- GHR dysfunction (GH resistance/insensitivity)
- Post-GHR signaling defect (STAT5b)
- *IGF1* gene defect
- ALS deficiency
- Hepatic insufficiency (eg, cirrhosis)
- Malnutrition
- Hypothyroidism
- Delayed puberty
- Poorly controlled diabetes mellitus
- Chronic illness
- Glucocorticoid therapy

Endocrinol Metab Clin N Am 41 (2012) 265–282

GH – IGF1 Axis

IPSS 2018

GH – IGF1 axis



Endocrinol Metab Clin N Am 36 (2007) 131–186

When to suspect Growth Hormone Deficiency (GHD)?

- Low growth velocity.
- Low IGF-1, IGF-BP₃
- Delayed bone age.
- Other causes were excluded
- ± Evidence of pituitary abnormality.

How to confirm GHD?

- Growth hormone stimulation testing:
 - Arginine,
 - Clonidine.
 - Glucagon
 - Levodopa
 - Insulin / hypoglycemia

Tests to Provoke Growth Hormone Secretion

Stimulus	Dosage	Times Samples Are Taken (min)	Comments
Exercise	Step climbing; exercise cycle for 10 min	0, 10, 20	Observe child closely when on the steps
Levodopa	< 15 kg: 125 mg 10-30 kg: 250 mg > 30 kg: 500 mg,	10, 60, 90	Nausea, rarely emesis
Clonidine	0.15 mgs/m ²	0, 30, 60, 90	Tiredness, postural hypotension
Arginine HCl (IV)	0.5 g/kg (max 30 g) 10% arginine HCl in 0.9% NaCl over 30 min	0, 15, 30, 45, 60	
Insulin (IV)	0.05-0.1 unit/kg	0, 15, 30, 60, 75, 90, 120	Hypoglycemia, requires close supervision
Glucagon (M)	0.03 mg/kg (max 1 mg)	0, 30, 60, 90, 120, 150, 180	Nausea, occasional emesis
GHRH (IV)	1 (g/kg)	0, 15, 30, 45, 60, 90, 120	Flushing, metallic taste

Tests should be performed after an overnight fast. Many investigators suggest that prepubertal children should be "primed" with gonadal steroids, eg, 5 mg Premarin orally the night before and the morning of the test or with 50 to 100 µg/d ethinyl estradiol for 3 consecutive days before testing or 100 ng depot testosterone 3 days before testing. This, of course, alters patient's steady state and performs the provocative test in a steroid-rich environment. Patients must be euthyroid at the time of testing.

Documentation of appropriate lowering of blood glucose is recommended. If GHD is suspected, the lower dosage of insulin is usually administered, especially in infants. D10W and glucagon should be available.

Copyright 2005 by Charles C Thomas, Publisher, LTD.

If GHD is confirmed

- MRI of brain (hypothalamus / pituitary)
- Start GH therapy.

IPS 2018

Reference

- Etiologies and Early Diagnosis of Short Stature and Growth Failure in Children and Adolescents. Alan D. Rogol, MD, PhD¹, and Gregory F. Hayden, MD
 - (J Pediatr 2014;164:S1-S14).
- **Kappy et al.** ■ *Pediatric Practice: Endocrinology 2010: 25*
- Endocrinol Metab Clin N Am 41 (2012) 265–282



Thank You